

SEQUENCE LISTING

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RECEIVED

TECH CENTER 1600/2900
<110> AMYLIN PHARMACEUTICALS, INC.
<120> NOVEL EXENDIN AGONIST COMPOUNDS
<130> 238/086 US (030639.0043.UTL2)
<140> 09/554,533
<141> 1998-11-13
<150> PCT/US98/24210
<151> 1998-11-13
<150> US 60/065,442
<151> 1997-11-14
<160> 74
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<211> 39
<212> PRT
<213> Heloderma horridum
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<221> AMIDATION
<222> (39)...(39)
<223> amidated Ser (Serinamide)
<400> 1
 His Ser Asp Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
             20
 Ser Gly Ala Pro Pro Pro Ser
         35
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<213> Heloderma suspectum
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<221> AMIDATION
<222> (39)...(39)
<223> amidated Ser (Serinamide)
<400> 2
 His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
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Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser 25 Ser Gly Ala Pro Pro Pro Ser 35 <210> 3 <211> 30 <212> PRT <213> Homo sapien <220> <221> AMIDATION <222> (30)...(30) <223> amidated Arg (Argininamide) <400> 3 His Ala Glu Gly Thr Phe Thr Ser Asp Val Ser Ser Tyr Leu Glu Gly 10 Gln Ala Ala Lys Glu Phe Ile Ala Trp Leu Val Lys Gly Arg 25 <210> 4 <211> 29 <212> PRT <213> Artificial Sequence <220> <223> artificially synthesized sequence of novel exendin agonist compound <220> <221> VARIANT <222> (1) ... (7) <223> Xaa in position 1 is His, Arg or Tyr; Xaa in position 2 is Ser, Gly, Ala or Thr; Xaa in position 3 is Asp or Glu; Xaa in position 5 is Ala or Thr; Xaa in position 6 is Ala, Phe, Tyr or naphthylalanine; Xaa in position 7 is Thr or Ser; <220> <221> VARIANT <222> (8)...(13) <223> Xaa in position 8 is Ala, Ser or Thr; Xaa in position 9 is Asp or Glu; Xaa in position 10 is Ala, Leu, Ile, Val, pentylglycine or Met; Xaa in position 11 is Ala or Ser; Xaa in position 12 is Ala or Lys; Xaa in position 13 is Ala or Gln; <220> <221> VARIANT

<223> Xaa in position 14 is Ala, Leu, Ile, pentylglycine, Val or

19 is Ala or Val; Xaa in position 20 is Ala or Arg;

Met; Xaa in position 15 is Ala or Glu; Xaa in position 16 is Ala or Glu; Xaa in position 17 is Ala or Glu; Xaa in position

<222> (14)...(20)

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<220>
<221> VARIANT
<222> (21) . . . (24)
<223> Xaa in position 21 is Ala or Leu; Xaa in position 22 is Phe,
     Tyr or naphthylalanine; Xaa in position 23 is Ile, Val, Leu,
     pentylglycine, tert-butylglycine or Met; Xaa in position 24
     is Ala, Glu or Asp;
<220>
<221> VARIANT
<222> (25) ... (28)
<223> Xaa in position 25 is Ala, Trp, Phe, Tyr or naphthylalanine;
     Xaa in position 26 is Ala or Leu; Xaa in position 27 is Ala
     or Lys; Xaa in position 28 is Ala or Asn;
<220>
<221> VARIANT
<222> (29)...(29)
<223> Xaa in position 29 is -OH; -NH2, Gly-Z2, Gly Gly-Z2, Gly Gly
     Xaa31-Z2; Gly Gly Xaa31 Ser-Z2; Gly Gly Xaa31 Ser Ser-Z2; Gly Gly
     Xaa31 Ser Ser Gly-Z2; Gly Gly Xaa31 Ser Ser Gly Ala-Z2; Gly Gly
     Xaa31 Ser Ser Gly Ala Xaa36-Z2,
<220>
<221> VARIANT
<222> (29) ... (29)
<223> Gly Gly Xaa31 Ser Ser Gly Ala Xaa36 Xaa37-Z2; or Gly Gly Xaa31
     Ser Ser Gly Ala Xaa36 Xaa37 Xaa38-Z2;
<220>
<221> VARIANT
<222> (29)...(29)
<223> where Xaa31, Xaa36, Xaa37 and Xaa38 are independently Pro,
     homoproline, 3Hyp, 4Hyp, thioproline, N-alkylglycine,
     N-alkylpentylglycine or N-alkylalanine; and Z2 is -OH or -NH2;
<220>
<221> VARIANT
<222> (3)...(28)
<223> provided that no more than three of Xaa in positions 3, 5, 6,
     8, 10, 11, 12, 13, 14, 15, 16, 17, 19, 20, 21, 24, 25, 26,
     27 and 28 are Ala.
<400> 4
10
20
                               25
<210> 5
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<213> Artificial Sequence
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<220>

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<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (30) ... (30)
<223> amidated Gly (Glycinamide)
<400> 5
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly
                                  25
<210> 6
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<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 6
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn
             20
<210> 7
<211> 28
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<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 7
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
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Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
20 25
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<210> 8
<211> 28
<212> PRT
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<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 8
His Ala Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                  5
                                      10
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 9
<211> 28
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<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 9
His Gly Glu Gly Ala Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
<210> 10
<211> 28
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
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compound

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<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 10
His Gly Glu Gly Thr Ala Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 11
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) . . . (28)
<223> amidated Asn (Asparaginamide)
<400> 11
His Gly Glu Gly Thr Phe Thr Ala Asp Leu Ser Lys Gln Leu Glu Glu
                  5
                                      10
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 12
<211> 28
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 12
His Gly Glu Gly Thr Phe Thr Ser Asp Ala Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
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<210> 13
<211> 28
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 13
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ala Lys Gln Leu Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 14
<211> 28
<212> PRT
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<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 14
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Ala Gln Leu Glu Glu
                  5
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
<210> 15
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<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
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<223> amidated Asn (Asparaginamide)
<400> 15
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Ala Leu Glu Glu
                  5
                                      10
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
<210> 16
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<223> artificially synthesized sequence of novel exendin agonist
      compound
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<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 16
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Ala Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 17
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<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 17
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Ala Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 18
<211> 28
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<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 18
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Ala
                                      10
 Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 19
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<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 19
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
 Ala Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
<210> 20
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) . . . (28)
<223> amidated Asn (Asparaginamide)
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<400> 20

His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu

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Glu Ala Ala Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 21
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 21
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                  5
                                      10
Glu Ala Val Ala Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 22
<211> 28
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 22
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Ala Phe Ile Glu Phe Leu Lys Asn
             20
<210> 23
<211> 28
<212> PRT
<213> Artificial Sequence
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<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 23
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Ala Phe Leu Lys Asn
             20
<210> 24
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 24
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                  5
                                      10
Glu Ala Val Arg Leu Phe Ile Glu Ala Leu Lys Asn
             20
<210> 25
<211> 28
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 25
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                                     10
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Glu Ala Val Arg Leu Phe Ile Glu Phe Ala Lys Asn 20 25

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<210> 26
<211> 28
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<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 26
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Ala Asn
             20
<210> 27
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Ala (Alaninamide)
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                  5
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Ala
             20
<210> 28
<211> 38
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
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compound

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<220>
<221> AMIDATION
<222> (38)...(38)
<223> amidated Pro (Prolinamide)
<400> 28
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
                                  25
 Ser Gly Ala Pro Pro Pro
         35
<210> 29
<211> 38
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (38)...(38)
<223> amidated Pro (Prolinamide)
<400> 29
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                  5
 Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro Ser
 Ser Gly Ala Pro Pro Pro
<210> 30
<211> 37
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (37) . . . (37)
<223> amidated Pro (Prolinamide)
```

<400> 30

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His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
             20
                                 25
 Ser Gly Ala Pro Pro
         35
<210> 31
<211> 37
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<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (37)...(37)
<223> amidated Pro (Prolinamide)
<400> 31
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro Ser
             20
                                                      30
 Ser Gly Ala Pro Pro
         35
<210> 32
<211> 36
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (36)...(36)
<223> amidated Pro (Prolinamide)
<400> 32
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
                                 25
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Ser Gly Ala Pro
         35
<210> 33
<211> 36
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence intended to act on
exendin agonist
<220>
<221> AMIDATION
<222> (36) ... (36)
<223> amidated Pro (Prolinamide)
<400> 33
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                  5
                                      10
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro Ser
                                  25
 Ser Gly Ala Pro
         35
<210> 34
<211> 35
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (35)...(35)
<223> amidated Ala (Alaninamide)
<400> 34
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
             20
                                 25
Ser Gly Ala
         35
<210> 35
<211> 35
<212> PRT
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<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (35)...(35)
<223> amidated Ala (Alaninamide)
<400> 35
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro Ser
 Ser Gly Ala
         35
<210> 36
<211> 34
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (34)...(34)
<223> Amidated Gly (Glycinamide)
<400> 36
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
                                 25
 Ser Gly
<210> 37
<211> 34
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
<220>
<221> AMIDATION
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<222> (34)...(34)
<223> amidated Gly (Glycinamide)
<400> 37
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro Ser
Ser Gly
<210> 38
<211> 33
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (33)...(33)
<223> amidated Ser (Serinamide)
<400> 38
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
                  5
                                      10
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
             20
                                 25
                                                      30
Ser
<210> 39
<211> 33
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (33)...(33)
<223> amidated Ser (Seinamide)
<400> 39
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro Ser
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20 25 30

Ser

<210> 40

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> artificially synthesized sequence of novel exendin agonist
 compound

<220>

<221> AMIDATION

<222> (32)...(32)

<223> amidated Ser (Serinamide)

<400> 40

His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu 1 5 10 15

Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser 20 25 30

<210> 41

<211> 32

<212> PRT

<213> Artificial Sequence

<220>

<223> artificially synthesized sequence of novel exendin agonist compound

<220>

<221> AMIDATION

<222> (32)...(32)

<223> amidated Ser (Serinamide)

<400> 41

His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu 1 5 10 15

Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro Ser 20 25 30

<210> 42

<211> 31

<212> PRT

<213> Artificial Sequence

<220>

<223> artificially synthesized sequence of novel exendin agonist compound

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<220>
<221> AMIDATION
<222> (31) . . . (31)
<223> amidated Pro (Prolinamide)
<400> 42
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
                  5
                                      10
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro
                                  25
             20
<210> 43
<211> 31
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (31) ... (31)
<223> amidated Pro (Prolinamide)
<400> 43
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro
             20
<210> 44
<211> 30
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (30)...(30)
<223> amidated Gly (Glycinamide)
<400> 44
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly
             20
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<210> 45
<211> 29
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<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (29)...(29)
<223> amidated Gly (Glycinamide)
<400> 45
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
                  5
                                      10
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly
             20
<210> 46
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<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (29) ... (29)
<223> amidated Gly (Glycinamide)
<400> 46
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly
<210> 47
<211> 38
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<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Xaa in positions 31, 36, 37 and 38 stands for thioproline.
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<220>
<221> AMIDATION
<222> (38)...(38)
<223> amidated tPro (Thioprolinamide)
<400> 47
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
                  5
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Xaa Ser
                                  25
Ser Gly Ala Xaa Xaa Xaa
         35
<210> 48
<211> 38
<212> PRT
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<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Xaa in positions 36, 37 and 38 stands for thioproline.
<220>
<221> AMIDATION
<222> (38) . . . (38)
<223> amidated tPro (Thioprolinamide)
<400> 48
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
Ser Gly Ala Xaa Xaa Xaa
         35
<210> 49
<211> 37
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<223> Ala in position 31 is N-methyl ala.
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<220>
<221> AMIDATION
<222> (37) ... (37)
<223> amidated Pro (Prolinamide)
<400> 49
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Ala Ser
             20
 Ser Gly Ala Pro Pro
         35
<210> 50
<211> 37
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Ala in positions 31, 36 and 37 is N-methyl ala.
<220>
<221> AMIDATION
<222> (37)...(37)
<223> amidated N-methyl ala (N-methyl alaninamide)
<400> 50
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
                  5
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Ala Ser
                                 25
 Ser Gly Ala Ala Ala
         35
<210> 51
<211> 37
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Xaa in positions 31, 36 and 37 stands for homoproline.
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<220>

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<221> AMIDATION
<222> (37) . . . (37)
<223> amidated hPro (Homoprolinamide)
<400> 51
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
                  5
                                      10
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Xaa Ser
 Ser Gly Ala Xaa Xaa
         35
<210> 52
<211> 36
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Xaa in positions 31 and 36 stands for homoproline.
<220>
<221> AMIDATION
<222> (36) ... (36)
<223> amidated hPro (Homoprolinamide)
<400> 52
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Xaa Ser
                                  25
                                                       30
 Ser Gly Ala Xaa
         35
<210> 53
<211> 35
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (35) ... (35)
<223> amidated Ala (Alaninamide)
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<400> 53
Arg Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
                  5
                                      10
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Pro Ser
                                  25
 Ser Gly Ala
         35
<210> 54
<211> 30
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (30)...(30)
<223> amidated Gly (Glycinamide)
<400> 54
His Gly Asp Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly
             20
                                 25
<210> 55
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Xaa in position 6 stands for naphthylalanine.
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 55
His Gly Glu Gly Thr Xaa Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
```

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<210> 56
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 56
His Gly Glu Gly Thr Phe Ser Ser Asp Leu Ser Lys Gln Met Glu Glu
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn
             20
<210> 57
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
His Gly Glu Gly Thr Phe Ser Thr Asp Leu Ser Lys Gln Met Glu Glu
                  5
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn
             20
<210> 58
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
```

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<223> amidated Asn (Asparaginamide)
<400> 58
 His Gly Glu Gly Thr Phe Thr Ser Glu Leu Ser Lys Gln Met Ala Glu
                  5
 Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn
<210> 59
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Xaa in position 10 stands for pentylglycine.
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 59
 His Gly Glu Gly Thr Phe Thr Ser Asp Xaa Ser Lys Gln Leu Glu Glu
                  5
 Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 60
<211> 28
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
     compound
<220>
<223> Xaa in position 22 stands for naphthylalanine.
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 60
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                                     10
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Glu Ala Val Arg Leu Xaa Ile Glu Phe Leu Lys Asn 20 25

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<210> 61
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Xaa in position 23 stands for tert-butylglycine.
<220>
<221> AMIDATION
<222> (28)...(28)
<223> amidated Asn (Asparaginamide)
<400> 61
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu
Glu Ala Val Arg Leu Phe Xaa Glu Trp Leu Lys Asn
             20
                                  25
<210> 62
<211> 28
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (28) ... (28)
<223> amidated Asn (Asparaginamide)
<400> 62
His Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                                      10
Glu Ala Val Arg Leu Phe Ile Asp Phe Leu Lys Asn
             20
<210> 63
<211> 33
<212> PRT
<213> Artificial Sequence
```

<220>

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<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (33)...(33)
<223> amidated Ser (Serinamide)
<400> 63
His Gly Glu Gly Thr Phe Thr Ser Asp Ala Ser Lys Gln Leu Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly Pro Ser
                                  25
 Ser
<210> 64
<211> 29
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<221> AMIDATION
<222> (29)...(29)
<223> amidated Gly (Glycinamide)
<400> 64
His Gly Glu Gly Thr Phe Thr Ser Asp Ala Ser Lys Gln Met Glu Glu
Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly
             20
<210> 65
<211> 37
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Xaa in positions 31, 36 and 37 stand for homoproline.
<220>
<221> AMIDATION
<222> (37) . . . (37)
<223> amidated hPro (homoprolinamide)
```

<400> 65

His Gly Glu Gly Thr Phe Thr Ser Asp Ala Ser Lys Gln Met Glu Glu 1 5 10 15

Glu Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly Xaa Ser 20 25 30

Ser Gly Ala Xaa Xaa 35

<210> 66

<211> 28

<212> PRT

<213> Artificial Sequence

<220>

<223> artificially synthesized sequence of novel exendin agonist
 compound

<220>

<221> VARIANT

<222> (1) . . . (6)

<223> Xaa in position 1 is His, Arg, Tyr or 4-imidazopropionyl; Xaa in position 2 is Ser, Gly, Ala or Thr; Xaa in position 3 is Asp or Glu; Xaa in position 5 is Ala or Thr; Xaa in position 6 is Ala, Phe, Tyr or naphthylalanine;

<220>

<221> VARIANT

<222> (7)...(12)

<223 > Xaa in position 7 is Thr or Ser; Xaa in position 8 is Ala, Ser or Thr; Xaa in position 9 is Asp or Glu; Xaa in position 10 Ala, Leu, Ile, Val, pentylglycine or Met; Xaa in position 11 is Ala or Ser; Xaa in position 12 is Ala or Lys;

<220>

<221> VARIANT

<222> (13)...(19)

<223> Xaa in position 13 is Ala or Gln; Xaa in position 14 is Ala, Leu, Ile, pentylglycine, Val or Met; Xaa in position 15 is Ala or Glu; Xaa in position 16 is Ala or Glu; Xaa in position 17 is Ala or Glu; Xaa in position 19 is Ala or Val;

<220>

<221> VARIANT

<222> (20)...(22)

<223> Xaa in position 20 is Ala or Arg; Xaa in position 21 is Ala, Leu or Lys-NH 6 -R where R is Lys, Arg, C_1 - C_{10} straight chain or branched alkanoyl or cycloalkylalkanoyl; Xaa in position 22 is Phe, Tyr or naphthylalanine;

<220>

<221> VARIANT

<222> (23)...(26)

<223> Xaa in position 23 is Ile, Val, Leu, pentylglycine, tertbutylglycine or Met; Xaa in position 24 is Ala, Glu or Asp; Xaa in position 25 is Ala, Trp, Phe, Tyr or naphthylalanine; Xaa in position 26 is Ala or Leu;

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<220>
<221> VARIANT
<222> (27) . . . (27)
<223> Xaa in position 27 is Lys Asn, Asn Lys, Lys-NH<sup>e</sup>-R Asn, Asn
       Lys-NH<sup>c</sup>-R, Lys-NH<sup>c</sup>-R Ala, Ala Lys-NH<sup>c</sup>-R where R is Lys, Arg,
       C1-C10 straight chain or branched alkanoyl or cycloalkyl-
       alkanoyl;
<220>
<221> VARIANT
<222> (28)...(28)
<223> Xaa in position 28 is -OH; -NH<sub>2</sub>, Gly-Z<sub>2</sub>, Gly Gly-Z<sub>2</sub>,
      Gly Gly Xaa31-Z2, Gly Gly Xaa31 Ser-Z2, Gly Gly Xaa31 Ser
       Ser-Z<sub>2</sub>, Gly Gly Xaa<sub>31</sub> Ser Ser Gly-Z<sub>2</sub> Gly Gly Xaa<sub>31</sub> Ser
      Ser Gly Ala-Z<sub>2</sub>, Gly Gly Xaa<sub>31</sub> Ser Ser Gly Ala Xaa<sub>36</sub>-Z<sub>2</sub>,
<220>
<221> VARIANT
<222> (28)...(28)
<223> Xaa in position 28 is Gly Gly Xaa31 Ser Ser Gly Ala
      Xaa36 Xaa37-Z2; or Gly Gly Xaa31 Ser Ser Gly Ala Xaa36
      Xaa37 Xaa38-Z2;
<220>
<221> VARIANT
<222> (28)...(28)
<223> where Xaa_{31}, Xaa_{36}, Xaa_{37} and Xaa_{38} are independently selected
      from the group consisting of Pro, homoproline, 3Hyp, 4Hyp,
       thioproline, N-alkylglycine, N-alkylpentylglycine or
      N-alkylalanine; and Z_2 is -OH or -NH_2;
<220>
<221> VARIANT
<222> (3) ... (26)
<223> provided that no more than three of Xaa in positions 3,
       5, 6, 8, 10, 11, 12, 13, 14, 15, 16, 17, 19, 20, 21, 24
      25 and 26 are Ala.
<400> 66
 10
Xaa Ala Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
              20
                                     25
<210> 67
<211> 27
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
```

<220>

compound

```
<223> Gly in position 1 is 4-imidazolylpropionyl-Gly.
      Lys in position 26 is Lys-NH<sup>6</sup>octanoyl.
<220>
<221> AMIDATION
<222> (27)...(27)
<223> amidated Asn (Asparaginamide)
<400> 67
 Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu Glu
                   5
 Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn
              20
<210> 68
<211> 27
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Gly in position 1 is 4-imidazolylpropionyl-Gly.
      Lys in position 26 is Lys-NH<sup>E</sup>octanoyl.
<220>
<221> AMIDATION
<222> (27) ... (27)
<223> amidated Asn (Asparaginamide)
<400> 68
Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu
                   5
                                       10
Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn
             20
<210> 69
<211> 29
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Gly in position 1 is 4-imidazolylpropionyl-Gly.
      Lys in position 26 is Lys-NH<sup>6</sup>octanoyl.
<220>
<221> AMIDATION
```

```
<222> (29) . . . (29)
<223> amidated Gly (Glycinamide)
<400> 69
 Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu Glu
 Ala Val Arg Leu Phe Ile Glu Trp Leu Lys Asn Gly Gly
             20
                                   25
<210> 70
<211> 29
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Gly in position 1 is 4-imidazolylpropionyl-Gly.
      Lys in position 26 is Lys-NH<sup>c</sup>octanoyl.
<220>
<221> AMIDATION
<222> (29) ... (29)
<223> amidated Gly (Glycinamide)
<400> 70
 Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu Glu
                                       10
 Ala Val Arg Leu Phe Ile Glu Phe Leu Lys Asn Gly Gly
             20
<210> 71
<211> 27
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<223> Gly in position 1 is 4-imidazolylpropionyl-Gly.
      Lys in position 27 is Lys-NH<sup>6</sup>octanoyl.
<220>
<221> AMIDATION
<222> (27) . . . (27)
<223> amidated Lys (Lysinamide)
<400> 71
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Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu Glu
 Ala Val Arg Leu Phe Ile Glu Trp Leu Asn Lys
              20
<210> 72
<211> 27
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Gly in position 1 is 4-imidazolylpropionyl-Gly.
      Lys in position 27 is Lys-NH<sup>6</sup>octanoyl.
<220>
<221> AMIDATION
<222> (27)...(27)
<223> amidated Lys (Lysinamide)
<400> 72
 Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu Glu
 Ala Val Arg Leu Phe Ile Glu Phe Leu Asn Lys
             20
<210> 73
<211> 29
<212> PRT
<213> Artificial Sequence
<220>
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Gly in position 1 is 4-imidazolylpropionyl-Gly.
      Lys in position 27 is Lys-NH<sup>6</sup>octanoyl.
<220>
<221> AMIDATION
<222> (29)...(29)
<223> amidated Gly (Gylcinamide)
<400> 73
Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Met Glu Glu Glu
Ala Val Arg Leu Phe Ile Glu Trp Leu Asn Lys Gly Gly
             20
                                  25
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<210> 74
<211> 29
<212> PRT
<213> Artificial Sequence
<223> artificially synthesized sequence of novel exendin agonist
      compound
<220>
<223> Gly in position 1 is 4-imidazolylpropionyl-Gly.
      Lys in position 27 is Lys-NH^{\epsilon}octanoyl.
<220>
<221> AMIDATION
<222> (29)...(29)
<223> amidated Gly (Glycinamide)
<400> 74
Gly Glu Gly Thr Phe Thr Ser Asp Leu Ser Lys Gln Leu Glu Glu Glu
                                      10
Ala Val Arg Leu Phe Ile Glu Phe Leu Asn Lys Gly Gly
```

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